

TABLE A2.—*Smoking and chronic obstructive pulmonary disease symptoms*<sup>1</sup>—percent prevalence (cont.)

(Numbers in parentheses represent total number of individuals in particular smoking group)

SM = Smokers.

NS = Nonsmokers.

EX = Ex-smokers.

Author, year, country, reference	Number and type of population	Cough	Sputum production	Breathlessness or dyspnea	Chest illnesses	Other	Comments
Wynder et al., 1965 U.S.A. (238).	315 male patients in New York City and 315 male patients in California.	<i>New York City</i> NS . . . . .14.0 (44) Pipe, cigar 33.0 (54) Cigarettes: 1-10 . . .45.0 (44) 10-20 . . .46.0 (88) >20 . . .67.0 (85) <i>California</i> NS . . . . .22.0 (69) Pipe, cigar 30.0 (32) Cigarettes: 1-10 . . .45.0 (54) 10-20 . . .74.0 (91) >20 . . .74.0 (69)					
Freour et al., 1966 France (92).	1,055 randomly chosen males in Bordeaux 30-70 years of age.					<i>Clinical signs of bronchitis and respiratory insufficiency</i> NS . . . . .25.4 (45) SM . . . . .54.4 (478)	
Haynes, et al., 1966 U.S.A. (108).	179 male preparatory school students 14-19 years of age.					<i>Average number of severe respiratory illnesses per 10 students (adjusted for age)</i> NS . . . . .0.36 All smokers 2.30 Heavy SM 3.34	Heavy smoker— more than 10 ciga- rettes per day.

TABLE A2.—*Smoking and chronic obstructive pulmonary disease symptoms<sup>1</sup>—percent prevalence (cont.)*

(Numbers in parentheses represent total number of individuals in particular smoking group)

SM = Smokers. NS = Nonsmokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	Cough	Sputum production	Breathlessness or dyspnea	Chest illnesses	Other	Comments
Densen et al., 1967, U.S.A. (68).	5,313 male and 7,291 female postal and transit workers.	<i>Postal</i> NS ..... 7.0 (903) Pipe, cigar 12.4 (628) Cigarettes only .... 27.0 (2,687) <i>Transit</i> NS ..... 6.4 (1,012) Pipe, cigar 10.5 (765) Cigarettes only .... 23.5 (3,745)	<i>Postal</i> 13.1 17.4  28.9 <i>Transit</i> 9.5 14.1 23.7	<i>Postal</i> 19.8 24.8  31.7 <i>Transit</i> 11.7 14.2 21.9			Dyspnea represented by Grade II only.
Higgins et al., 1968, U.S.A. (118).	926 white male resi- dents of Marion County, West Virginia, 26-69 years of age.	NS ..... 15.4 (162) SM ..... 47.2 (513) EX ..... 19.3 (144)	NS ..... 31.1 SM ..... 46.2 EX ..... 28.5	NS ..... 5.0 SM ..... 10.7 EX ..... 16.8			
Holland and Elliott, 1968, England (121).	9,786 male and female school children.	<i>Males</i> NS ..... 3.8 (1,900) SM ..... 6.3 (1,098) EX ..... 2.9 (1,782) <1 cigarette/day ..... 1-2 ..... 3-4 ..... >5 .....	<i>Females</i> 3.2 (3,137) 6.3 (554) 4.3 (1,151) 5.8 (876) 6.5 (417) 6.6 (124) 9.9 (142)	<i>Males</i> 2.4 6.1 3.9    18.3 <i>Females</i> 2.1 8.3 4.2 5.8 8.4 8.1			

TABLE A2.—*Smoking and chronic obstructive pulmonary disease symptoms<sup>1</sup>—percent prevalence (cont.)*

(Numbers in parentheses represent total number of individuals in particular smoking group)

SM = Smokers.

NS = Nonsmokers.

EX = Ex-smokers.

Author, year, country, reference	Number and type of population	Cough	Sputum production	Breathlessness or dyspnea	Chest illnesses	Other	Comments
Gandevia 1969 Australia (93).	762 male and 1,304 female patients from 13 general practices in all parts of Australia.	<i>Males</i> NS .....10.3 (234) SM .....51.3 (528) <i>Females</i> NS .....10.5 (857) SM .....37.4 (447)					Productive cough upon request.
Rimington 1969 England (193).	41,729 male and 22,295 female persons participating in mass miniature radiography screening.					<i>Age-adjusted total prevalence of chronic bronchitis</i> <i>Males</i> NS ..... 5.1 (9,055) EX ..... 9.8 (6,510) Pipe ..... 9.0 (2,921) Cigarettes . . (23,243) 1- 9 ..... 9.1 10-19 ..... 15.0 >20 ..... 20.6 <i>Females</i> NS ..... 3.4 (12,351) EX ..... 3.8 (959) Pipe ..... 0.0 Cigarettes . . (8,985) 1- 9 ... 5.1 10-19 ... 10.6 >20 ..... 18.5	Cigarette dosage gradient significant to $p < 0.001$ .
Wilhelmsen et al., 1969, Sweden (231).	313 males 50-54 years of age randomly sampled from population of Göteborg.					<i>Chronic bronchitis</i> NS ..... 1.0 (88) EX ..... 3.0 (67) 1-14 grams/ day ... 5.0 (94) >15 ..... 17.0 (64)	

TABLE A2.—*Smoking and chronic obstructive pulmonary disease symptoms<sup>1</sup>—percent prevalence (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 SM = Smokers. NS = Nonsmokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	Cough	Sputum production	Breathlessness or dyspnea	Chest illnesses	Other	Comments
Lambert and Reid, 1970, England (146).	9,975 male and female responders to a postal survey (4,688 males and 5,287 females 35-69 years of age).	<i>Persistent cough and phlegm</i>					
		<i>Males</i>					
			<i>Age</i>	<i>Age</i>	<i>Age</i>		
			35-45	45-55	55-65	65-69	
		NS . . . . .	7 (227)	6 (200)	11 (171)	7 (61)	
		EX . . . . .	7 (303)	11 (358)	15 (335)	18 (148)	
		<20 . . . . .	15 (521)	22 (488)	30 (490)	37 (139)	
		20 . . . . .	23 (191)	28 (204)	32 (149)	38 (37)	
		>20 . . . . .	27 (148)	28 (136)	42 (121)	25 (12)	
		<i>Females</i>					
		NS . . . . .	3 (500)	4 (637)	5 (925)	6 (21)	
		EX . . . . .	3 (127)	8 (128)	7 (94)	7 (41)	
Lefcoe and Wonnacott, 1970, Canada (151).	310 male physicians in London and Ontario, 25-74 years of age.	<20 . . . . .	9 (602)	13 (472)	16 (306)	11 (65)	
		20 . . . . .	16 (128)	27 (122)	31 (77)	14 (7)	
		>20 . . . . .	23 (22)	26 (39)	43 (7)	.. (1)	
						<i>Age-standardized rates of chronic respiratory disease</i>	Excluded from ex-smokers are those
						NS . . . . . 1.0 (88)	cigarette
						EX . . . . . 5.0 (61)	smokers who
						SM . . . . . 34.0 (101)	now smoke
						Pipe, cigar . . 12.0 (33)	pipes or cigars.

<sup>1</sup> Data collected by either direct interview, questionnaire, review of medical records and/or medical examination.

TABLE A2a.—*Smoking and chronic obstructive pulmonary disease symptoms*<sup>1</sup>—percent prevalence

(Numbers in parentheses represent total number of individuals in particular smoking group)

SM = Smokers. NS = Nonsmokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of popuation	Cough		Bronchitis			Comments		
Cederlof et al., 1966, Sweden (46).	9,319 twin pairs registered in Sweden of 12,889 available.		<i>Observed/ expected cases</i>	<i>Hypermorbidity ratio</i>	<i>Observed expected cases</i>	<i>Hypermorbidity ratio</i>	Explanation of analyses for respiratory symptom prevalence:	All ex-smokers included with smokers.	
		Group A:							
		Males	393/151.9	2.6	157/50.8	3.1	Group A analysis—using each firstborn twin as one group in an unmatched relationship to each secondborn twin.	MZ—monozygotic pairs	
		Females	136/ 49.4	2.8	43/11.2	3.8	Group B analysis—using each twin set as matched pair.	DZ—dizygotic pairs	
		Group B SM/NS:						Author concludes that since hypermorbidity for smoking persists in smoking-discordant MZ population, a casual relationship of smoking and broncho- pulmonary symptoms is supported.	
		MZ Males	14.6/7.7	1.9	6.6/ 1.1	6.0 (274)			
		Females	13.6/7.6	1.8	3.0/ 2.3	1.33 (264)			
		DZ Males	12.3/5.5	2.25	4.5/ 1.8	2.54 (733)			
	Females	14.5/5.7	2.57	5.5/ 1.8	3.0 (653)	All comparisons in Groups A and B are between smoking- discordant pairs.			
Cederlof et al., 1969, U.S.A. (45).	4,379 twin pairs (all U.S. veterans) in U.S. National Academy of Sciences Twin Registry (of 9,000 avail- able).	<i>Prevalence of respiratory symptoms</i>							
		Group A:							No ex-smokers included in Group B analysis.
		NS	4.3	4.3		1.6	Group A—as above.		The authors conclude that the data indicate a strong probability of a causal connection with smoking. Even these symptoms, however, seem to be influenced by genetic factors.
		1-10	6.4	6.4		2.7	Group B—as above.		
		11-30	15.3	15.3		8.0			
		>31	27.7	27.7		16.8			
		Pipe, cigar	7.1	7.1		2.7			
		Group B:		NS	SM	NS	SM		
		MZ	2.4	5.4	1.8	4.8			
		DZ	2.0	9.8	1.6	9.1			

<sup>1</sup> Data collected by either direct interview, questionnaire, review of medical records and/or medical examination.

TABLE A3.—*Smoking and ventilatory function*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	MBC	EFR	FEV	VC	Miscellaneous	Comments
Chivers, 1959, England (52).	463 male employees of alkaline industry plant.	Cigarettes/day: 0-5 ..... 6-20 ..... >20 .....	64'' †97 (28) 89 (50) 63 (6)	Height-in-inches 66'' 91 (35) 88 (75) 88.5 (9)	68'' 108 (31) 101 (112) 92.5 (9)	70'' 101 (21) 109 (75) 113 (12)	†Mean EFR in liters per minute. Regression analysis of data revealed a significant re- lationship between smoking and de- creasing function.
Higgins et al., 1959, England (116).	773 males in various occupations (25-34 and 55-64 years of age).	25-34 NS 145 (56) EX 143 (31) (25-34 and 1-14 grams .140 (193) >15 grams .133 (89)	55-64 101 (29) 89 (62) 87 (157) 80 (136)				FEV <sub>0.75</sub> expressed as mean indirect MBC.
Wilson et al., 1960 U.S.A. (232).	28 male residents of Dallas, Texas, former rural dwellers; matched for body surface, age, and height.					RV/TLC NS ..... 5.59 (14) NS ..... 21.1 SM ..... 4.44 (14) SM ..... 27.01	

TABLE A3.—*Smoking and ventilatory function (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	MBC	EFR	FEV	VC	Miscellaneous	Comments				
Ashford et al., 1961, Scotland (11).	4,014 male coal workers at 3 Scottish collieries.			<i>FEV<sub>1.0</sub></i>			Data represent results after correction for sitting height. SM includes pipe smoker. Data on ex-smoker not included. FEV <sub>1.0</sub> found significant; lower for SM than NS.				
				Age:	<i>NS</i>			<i>SM</i>			
				<21-30	4.09 (103)			3.96 (280)			
				21-30	3.86 (182)			3.77 (555)			
				31-40	3.44 (138)			3.88 (777)			
				41-50	3.04 (110)			2.96 (755)			
				51-60	2.71 (102)			2.56 (610)			
	>60 ...	2.38 (42)	2.21 (237)								
Fletcher and Tinker, 1961, England (85).	363 male London transport employees.		<i>Mean peak EFR</i>								
			NS .....	570 (30)							
			1-14 grams	537 (156)							
			>15 grams	528 (116)							
		EX .....	555 (61)								
Franklin and Lowell, 1961, U.S.A. (87).	213 male factory workers 40-60 years of age.			<i>FEV<sub>1.0</sub></i> <i>FEV<sub>0.25</sub></i> <i>FEV<sub>0.50</sub></i>			Heavy smoker represents an amount equal to or more than 30 pack years.				
				Heavy	2,670			3,011	2,710	Light ..	3,703 (59)
				Light	2,489			2,656	2,284	Heavy	3,578 (104)

TABLE A3.—*Smoking and ventilatory function (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	MBC	EFR	FEV	VC	Miscellaneous	Comments
Balchum et al., 1962, U.S.A. (24).	1,451 male employees in California light indus- try.		<i>MMEFR</i> NS ..... 15.5 (38) Pack/year: <1 ..... 15.0 (257) 1-9 ..... 10.0 (263) 10-19 ... 10.0 (303) 20-29 ... 19.0 (236) 30-39 ... 33.0 (144) 40-49 ... 38.0 (92) 50-59 ... 55.0 (29) >60 ..... 71.0 (24)	7.8 (19)  8.0 6.0 12.0 24.0 26.0 40.0 45.0 62.0			Data for: MMEFR given as percent of individuals with a value of <500 L/M; FEV <sub>1.0</sub> given as percent of individuals with value of <70 percent of expected.
Goldsmith et al., 1962, U.S.A. (95).	3,311 active or retired longshore- men.		<i>MEFR</i> NS ..... 313.63 (250) Pipe, cigar 299.26 (125) EX ..... 295.23 (102) Cigarettes/day: ≤20 ..... 309.73 (144) 20-40 ... 303.44 (346) ≥40 ..... 307.63 (57)	<i>FEV<sub>1.0</sub></i> 2.99 2.80 2.84  2.89 2.91 2.90			Authors concluded that cigarette smoke was found to have a slight effect on pulmonary function.
Martt, 1962, U.S.A. (161).	73 healthy medical per- sonnel with- out signifi- cant age difference between smokers and nonsmokers.				<i>D<sub>L</sub>CO</i> NS ..... 33.10 (30) SM <5 years .. 28.40 (8) 5-10 years ... 28.20 (10) >10 years ... 24.90 (25)		Smokers defined as those smoking >20 cigarettes/ day for varying periods.



TABLE A3.—*Smoking and ventilatory function (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	MBC	EFR	FEV	VC	Miscellaneous	Comments		
Revotskie et al., 1962, U.S.A. (192).	1,130 male and 1,813 female residents in Framing- ham partici- pating in the pro- spective study.			<i>FEV<sub>1.0</sub></i> <i>Males Females</i>			Data presented in terms of ratio of observed to predicted values.		
				NS	.....0.98 (55) 0.98 (255)				
				Cigarettes/day:					
				1-10	.0.97 (90) 0.99 (92)				
				10-29	.0.91 (163) 0.93 (157)				
				>30	..0.90 (81) 0.91 (22)				
Krumholz et al., 1964, U.S.A. (140).	18 physicians 24-37 years of age.		<i>MEFR</i> NS ..... 580 (9) SM ..... 1590 (9)			<i>Mean D<sub>L</sub></i> NS SM			
						Rest	.....36 31		
						Exercise:			
						2 minutes	..50 41		
						4 minutes	..50 43		
						3 minutes post exercise	39 135		
Zwi et al., 1964, U.S.A. (241).	20 medical students or graduate physicians.	NS ... 187 (10) SM ... 193 (10)	<i>MMEFR</i> 4.34 15.09	5.77 15.53		Authors found a significant difference between SM and NS for RV/TLC, compliance, and non- elastic resistance.			
Coates et al., 1965, U.S.A. (53).	1,342 male and 242 female post office employees >40 years of age.			<i>FEV<sub>1.0</sub></i> <i>Age: NS &gt;25 cig/day</i>		<i>Timed VC</i> <i>NS &gt;25/day</i>		<i>FEV<sub>1.0</sub>/VC</i> <i>NS &gt;25/day</i>	
				40-44	12.99 (186) 2.85 (69)	3.89 3.85	3 0.77 0.74		
				45-49	12.95 (170) 2.64 (42)	3.92 3.83	3 0.74 0.70		
				50-54	12.75 (115) 2.62 (22)	3.71 3.74	2 0.74 0.70		
				55-59	12.64 (64) 2.44 (18)	3.54 3.61	5 0.74 0.68		
				60-64	12.35 (53) 2.30 (8)	3.30 3.33	1 0.72 0.70		

TABLE A3.—*Smoking and ventilatory function (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	MBC	EFR	FEV	VC	Miscellaneous	Comments
			PEFR†	FEV <sub>1.0</sub> ‡	Forced VC‡		
			Males Females	Males Females	Males Females		
Huhti, 1965, Finland (126).	653 male and 823 female residents of a rural region in Finland.	NS ..... EX ..... Cigarettes/day: 1-14 ..... 15-24 ..... >25 .....	569 (122) 410 (709) 551 (141) 403 (30) 518 (108) 537 (191) 517 (85)	3.46 2.42 3.39 2.32 3.17 2.74 3.30 3.08	4.40 3.18 4.51 3.19 4.40 3.53 4.51 4.26		Pipe and cigar smokers not included. † Difference between NS and >25/day is significant for 45-49, 60-64 age groups. ‡ Trend is not statistically significant.
Krumholz et al., 1965, U.S.A. (142).	20 male medical students or graduate physicians.					Pulmonary compliance NS ..... 0.241 (10) SM ..... 0.177 (10) Compliance/FRC NS ..... 0.054 SM ..... 0.042	Mean body surface area for 2 groups was not signifi- cantly different. Smokers are those with equal to or greater than 5 pack year history.
Rankin et al., 1965, U.S.A. (189).	125 males without a past history of respira- tory disease 20-63 years of age.	NS ... 118.1 (68) SM ... 111.7 (57)		FEV <sub>1.0</sub> NS ..... 106.6 SM ..... 102.7		D <sub>L</sub> NS ... 31.1 SM ... 25.9 D <sub>L</sub> /alveolar volume 6.22 4.96	NS includes pipe and cigar smokers and ex-smokers of greater than 1 pack year. D <sub>L</sub> values have been corrected for COHb.

TABLE A3.—*Smoking and ventilatory function (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	MBC	EFR	FEV	VC	Miscellaneous	Comments
Edelman et al., 1966, U.S.A. (73).	410 male community dwellers 20-103 years of age.	NS ..... 164 (152) Current cigarette smokers. . <sup>2</sup> 151 (118) EX ..... 157 (93) Pipe, cigar .. 167 (47)	7.89  7.36 8.09 8.20	<i>FEV</i> <sub>1.0</sub> 2.83  <sup>2</sup> 2.64 2.80 2.91	<i>Vital capacity</i> 4.93  <sup>3</sup> 4.74 4.77 5.08		Ex-smokers of cigarettes only. Difference signifi- cant between NS and current cigarette smokers at p<0.01.
Peters and Ferris, 1967, U.S.A. (182).	124 male college age students.	NS ..... <sup>2</sup> 10.28 (41) Moderate . 10.06 (54) Heavy ..... 9.64 (29) EX ..... 9.48 (10)	<i>MEFR</i>     	<i>FEV</i> <sub>1.0</sub> 4.63 4.59 4.43 4.74	<i>FEV</i> <sub>1.0</sub> / <i>VC</i> <sup>2</sup> 87.5 85.3 83.9 83.2		Heavy smoker refers to greater than or equal to 4 pack years. Moderate smoker includes pipe and cigar smokers. Difference between NS and heavy smoker is significant.
Higgins et al., 1968, U.S.A. (118).	926 white male residents of Marion County, West Virginia, 20-69 years of age.	NS ..... EX ..... Cigarette SM 1-14 ..... 15-24 ..... >25 .....	<i>FEV</i> <sub>1.0</sub> 3.64 (160) 3.25 (143) 3.48 (511) 3.67 (88) 3.57 (273) 3.30 (150)				

TABLE A3.—*Smoking and ventilatory function (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	MBC	EFR	FEV			VC			Miscellaneous	Comments
Sluis- Cremer and Siehel, 1968, South Africa (208).	533 white male factory workers over 35 years of age.	NS	35-44	45-54	>55	FEV <sub>1.0</sub>			1 cigarette = 1 gram. 1 ounce tobacco = 26 grams. 1 cigar = 2 to 5 grams. † Derived slopes found signifi- cantly different from 0.		
			553 (106)	527 (101)	444 (27)	35-44	45-54	>55			
			Grams/day:			3.70	3.22	2.76			
			1-14	557 (26)	519 (17)	410 (7)	3.64	3.31		2.24	
			15-24	532 (94)	446 (35)	401 (13)	3.66	2.94		2.28	
			>25	†528 (66)	†494 (31)	†380 (10)	3.54	3.05		†2.12	
Stanescu et al., 1968, Rumania (212).	87 male bus drivers; 27 aged 20-25, 60 aged 40-60, all without respiratory symptoms.		FEV <sub>1.0</sub>			Nitrogen gradient					
			Younger	Older	Younger	Older	Younger	Older			
			NS	4,470 (14)	3,310 (40)	5,125	4,290	1.53		2.49	
			SM	4,500 (13)	†3,200 (20)	†5,285	†4,290	†1.47		†3.77	
Densen et al., 1969, U.S.A. (69).	5,287 male postal and transit workers in New York City.	NS	FEV <sub>1.0</sub> Postal			Non-white			FEV expressed as standardized for specified postal and transit workers at age 45 and at sitting height of 35 inches. Includes mixed smokers.		
			White								
			All cigarette	3.29 (685)	3.05 (204)						
			<25 grams/day	3.11 (2,340)	2.94 (768)						
			≥25 grams/day	3.14 (1,292)	2.95 (599)						
				3.06 (1,038)	2.93 (161)						
			Transit								
			White			Non-white					
			All cigarette	3.39 (620)	3.08 (298)						
			<25 grams/day	3.11 (2,941)	2.99 (1,041)						
			≥25 grams/day	3.15 (1,929)	3.00 (891)						
				3.02 (1,011)	2.95 (149)						

TABLE A3.—*Smoking and ventilatory function (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference	Number and type of population	MBC	EFR	FEV	VC	Miscellaneous	Comments
Rankin et al., 1969, Australia (190).	60 male and 10 female patients with chronic alcoholism 26-66 years of age.			<i>FEV<sub>1.0</sub></i> NS ..... 497.5 (12) SM ..... 78.4 (58)			FEV expressed as percent of predicted value for age, sex, and height.
Wilhelmsen et al., 1969, Sweden (231).	313 male residents of Göteborg 50-54 years of age.		NS ..... EX ..... 1-14 grams/day ..... >15 grams/day .....	<i>PEFR</i> 525 (88) 539 (67) 521 (94) 492 (64)	<i>FEV<sub>1.0</sub></i> 3.77 3.69 3.62 3.39	VC 4.83 4.77 4.83 4.56	1963 values only.
Lefcoe and Wonna- cott, 1970, Canada (151).	310 male physicians of London, Ontario.		<i>MMFR</i> NS ..... 4.09 (88) Cigarette smokers. 3.64 (101) EX ..... 3.99 (61) Pipe, cigar 4.17 (33)	<i>FEV<sub>1.0</sub></i> 3.39 3.11 3.38 3.17			MMFR has been standardized for age and height.

TABLE A3.—*Smoking and ventilatory function (cont.)*  
 (Numbers in parentheses represent total number of individuals in particular smoking group)  
 NS = Nonsmokers. SM = Smokers. EX = Ex-smokers.

Author, year, country, reference		FEV	Miscellaneous	Comments
Lundman, 1966, Sweden (159).	37 MZ and 62 DZ twin pairs selected from Swedish Twin-Pair Registry.	<u>FEV<sub>1.0</sub></u> Significant differences between smoking discordant twin pairs found for: 1. Group A MZ males and females. 2. Group B DZ males. 3. Group A DZ males.	<u>N<sub>2</sub> washout gradient</u> Significant differences between smoking dis- cordant twin pairs found for: Group B DZ males.	MZ = monozygotic. DZ = dizygotic. The author concludes that the degree of ventilation as measured by N <sub>2</sub> washout was correlated with cigarette consumption. The FEV <sub>1.0</sub> was significantly lower for smokers and there was a correlation with cigarette consumption. Explanation of analyses for respiratory symptom prevalence: Group A analysis—using each firstborn twin as one group in an unmatched relationship to each secondborn twin. Group B analysis—using each twin set as matched pair. All comparisons in Group A and B are between smoking-discor- dant pairs.

<sup>1</sup> Not significant (difference or trend).

<sup>2</sup>  $p < 0.05$

<sup>3</sup>  $p < 0.01$

<sup>4</sup>  $p < 0.005$

<sup>5</sup>  $p < 0.001$

TABLE A4.—Glossary of terms used in tables and text on smoking and ventilatory function

Symbol	Term	Volume or rate	Definition
MBC.....	Maximal breathing capacity.	Liters.....	The maximal volume of gas that can be breathed in one minute.
MVV.....	Maximal voluntary ventilation.		
EFR.....	Expiratory flow rate.....	Liters/minute.....	Rate of flow for a specified portion of a forced expiration (MMEFR—rate of flow measured for middle half of FVC).
PEFR.....	Peak expiratory flow rate.		
MEFR.....	Maximal expiratory flow rate.		
MMEFR.....	Maximal midexpiratory flow rate.		
FEV <sub>t</sub> .....	Forced expiratory volume.	Liters.....	Volume expired within a specified time interval. (FEV <sub>1.0</sub> —volume expired in first second of expiration.)
VC.....	Vital capacity.....	Liters.....	Maximal volume of a gas that can be expelled from the lungs by forceful effort following a maximal expiration.
FVC.....	Forced vital capacity.		
FEV <sub>t</sub> /VC....	Forced expiratory volume/vital capacity.	Percent.....	Volume of forced expiration (in time specified) related to vital capacity.
D <sub>L</sub> .....	Pulmonary diffusing capacity.	ml/min/mmHg	The ability of a chosen gas to pass from the alveolus to within the pulmonary capillary.
N <sub>2</sub> washout...	Nitrogen washout gradient.	Exponential curve.	The stepwise pulmonary alveolar clearance of a gas. (Slope of curve depends upon the uniformity and adequacy of ventilation of all parts of the lung.) It may be done as a single—or multiple—breath procedure.
	Compliance.....	Liters/CMH <sub>2</sub> O.....	Volume change of the lung produced by a unit pressure change.
RV.....	Residual volume.....	Liters.....	Volume of gas remaining in the lungs at the end of a maximal expiration.
TLC.....	Total lung capacity.....	Liters.....	Volume of gas contained in the lungs at the end of a maximal inspiration.
FRC.....	Functional residual capacity.	Liters.....	Volume of gas remaining in the lungs at the resting expiratory level.
	Alveolar volume.....	Liters.....	Volume of gas contained in pulmonary alveoli.

SOURCE: Comroe, J. et al. (56)

TABLE A6.—*Epidemiological studies concerning the relationship of air pollution, social class, and smoking to chronic obstructive bronchopulmonary disease (COPD)*

Author, year, country, reference	Number and type of population	Results
Higgins, 1957, England (112).	301 males and 280 females living in 2 separate districts. (45-64 years of age.)	Male data only (170): (a) The frequency of recurrent chest illnesses was higher in the more polluted region but the prevalence of other respiratory symptoms and mean values were similar. (b) Significant difference observed in COPD mortality rate.
College of General Practitioners, 1961, England (55).	787 males and 782 females 45-64 years of age from medical doctors' case lists.	(a) Male urban inhabitants manifested almost twice the prevalence of chronic bronchitis as rural males; this difference could not be explained on the basis of smoking habits. (b) No significant urban/rural differences noted for PEF <sub>R</sub> . <sup>1</sup> (c) No significant urban/rural differences noted for COPD symptoms among females.
Ferris and Anderson, 1962, U.S.A. (81).	1,219 males and females living in 3 different areas of a New Hampshire town.	Following adjustment for differences in smoking habits, no significant differences in chronic bronchitis were observed among the 3 pollution areas.
Mork, 1962, U.S.A. (171).	339 male transport employees from London and Norway.	The excess prevalence of serious respiratory symptoms (dyspnea, wheezing) and PEF <sub>R</sub> dysfunction among London Transport employees was only partly eliminated after standardization for smoking, and the author suggests that this is due to differences in air pollution levels.
Schoettlin, 1962, U.S.A. (204).	2,622 males 45-75 years of age.	(a) No positive correlation found between chronic respiratory illness and city size. (b) A positive correlation was found between chronic respiratory illness and cigarette smoking (particularly duration).
Anderson et al., 1965, Canada (8).	778 residents of Berlin, N.H., and 918 residents of Chilliwack, Canada.	Berlin, New Hampshire, has higher SO <sub>2</sub> and particulate air pollution levels and the higher respiratory disease prevalence rates among its residents were not accounted for by age differences, but were accounted for after standardization for smoking habits (except that PEF <sub>R</sub> and FEV <sub>1.0</sub> dysfunction was more prevalent in New Hampshire, and the authors suggest that this difference reflects air pollution differences).
Holland and Reid, 1965, England (124).	676 male transport employees in London and rural England.	(a) London employees manifested a greater prevalence of COPD symptoms and PEF <sub>R</sub> dysfunction than did the rural employees. (b) Smoking habit differences alone were not sufficient to explain this difference in COPD manifestations. (c) Both groups manifested pulmonary dysfunction correlated with tobacco consumption.
Bates et al., 1966, Canada (27).	216 hospitalized veterans from various areas of Canada (all standardized for age, tobacco consumption, and occupation).	Winnipeg (cleanest of all areas in SO <sub>2</sub> and industrial dustfall) residents manifested decreased prevalence of chest illnesses, less severe grades of dyspnea, and less sputum volume produced when compared to residents of all other areas.



TABLE A6.—*Epidemiological studies concerning the relationship of air pollution, social class, and smoking to chronic obstructive bronchopulmonary disease (COPD) (cont.)*

Author, year, country, reference	Number and type of population	Results
Ashley, 1969, England (12).	Standardized mortality ratios for males (1958-63) for 53 boroughs with air pollution indexes.	Positive correlations: (a) Smoke concentration and bronchitis mortality. (b) SO <sub>2</sub> and smoke concentration and bronchitis mortality and social class. (c) Pollution and social class.
Holland et al., 1969, England (122).	10,971 children over 11 years of age in 4 areas.	Factors affecting prevalence of respiratory symptoms: (a) Smoking—highly significant association. (b) Area of residence (pollution)—significant association except for periods of cough and phlegm lasting more than 3 weeks. (c) Social class, age, sex—no association noted.
Winkelstein and Kantor, 1969, U.S.A. (233).	842 females over 25 years of age in various regions of Buffalo.	(a) The increased prevalence of respiratory symptoms could not be explained by social class differences. (b) No overall association noted between productive cough and air pollution.
Cooley and Reid, 1970, England (58).	10,887 children 6-10 years of age from contrasting urban and rural areas.	Illnesses considered included chronic cough, past bronchitis, blocked nose. (a) Every geographic area showed a clear gradient of increasing illness prevalence with decreasing social class. (b) Social classes I, II, and III showed no urban/rural gradient while IV and V showed a clear excess in frequency of chest illnesses among urban residents over rural residents.
Lambert and Reid, 1970, England (146).	9,975 males and females responding to questionnaire survey.	(a) The trend of increasing prevalence of bronchitic symptoms from rural to urban respondents was not negated by adjustment for smoking differences. (b) After adjustment for age and smoking habits, male respondents manifested a clear correlation of persistent cough and phlegm prevalence with increasing air pollution. Correlation was not as striking in females. (c) Although the proportionate rise in symptom prevalence increased with air pollution similarly in each smoking group, the absolute differences in morbidity risk increased with increased cigarette consumption, suggesting synergistic influences of cigarette smoking and air pollution. (d) In the absence of cigarette smoking, the correlation between the prevalence of persistent cough and phlegm and air pollution was slight.

<sup>1</sup> See Glossary of Terms: Bronchopulmonary table A4.

TABLE A7.—*Epidemiological studies concerning the relationship of occupational exposure and smoking to chronic obstructive bronchopulmonary disease*

Author, year, country, reference	Number and type of population	Results
Higgins et al., 1956, England (119).	185 males (84 nonminers, 101 miners) without pneumoconiosis.	Miners showed increased symptom prevalence (breathlessness, cough, sputum). Miners showed increased prevalence of chronic bronchitis. Miners showed decreased MBC. <sup>1</sup> Differences in smoking between the two groups did not account for above differences.
Phillips et al., 1956, U.S.A. (185).	1,274 males factory employees (coke and electrolytic process).	None of the industrial environments were associated with an increased prevalence of chronic cough. Cigarette smoking and age were directly correlated with increased prevalence of chronic cough.
Higgins et al., 1959, England (116).	325 males 25-34 years of age and 401 males 55-64 years of age in various occupations.	Miners as compared to workers in non-dusty occupations: 25-34 years of age—significantly increased prevalence of chronic bronchitis and MBC abnormalities. 55-64 years of age—less significantly increased prevalence of chronic bronchitis and MBC abnormalities than in 25-34 years of age group. No smoking information available.
Chivers, 1959, England (52).	463 males in non-dusty and dusty occupations (lime and soda ash exposure).	No significant differences in PEFR <sup>1</sup> between dusty and non-dusty groups. Cigarette smoking (especially in those >40 years of age) was associated with decreased PEFR values.
Higgins and Cochrane, 1961, England (115).	300 male miners and 300 male nonminers 35-64 years of age.	Miners showed increased prevalence of symptoms and decreased MBC values which remained even after standardization for smoking habits. Total dust exposure was not directly correlated with these findings. Wives of miners showed similar symptom and test changes as compared with wives of nonminers.
Brinkman and Coates, 1962, U.S.A. (42).	1,317 males 40-65 years of age with various silica exposure histories.	Increased silica exposure was associated with an increased prevalence of chronic bronchitis. Highest prevalence of chronic bronchitis was noted in the non-exposed group; and this group was noted to have the highest number of smokers and highest consumption.
Hyatt et al., 1964, U.S.A. (128).	267 male miners and ex-miners 45-55 years of age.	Increased history of underground work was associated with an increased bronchopulmonary symptom prevalence and decreased pulmonary function values. The impairment of pulmonary function associated with underground work was separate from effect of smoking; but smoking and underground work did show additive effects.
Elwood et al., 1965, Ireland (77).	2,528 male and female flax workers over 35 years of age.	Preparing room workers who manifested byssinosis symptoms also showed an increased prevalence of chronic bronchitis independent of age or smoking when compared with non-preparing room workers. Female workers manifested a significant association between byssinosis symptoms and smoking while male workers did not.
Sluis-Cremer et al., 1967, South Africa (209).	827 miners and nonminers over 35 years of age.	Those smokers exposed to gold mine dust manifested more symptoms of COPD <sup>1</sup> than did non-dust exposed smokers, while prevalence of symptoms, among nonsmokers, was similar for the two groups.

TABLE A7.—*Epidemiological studies concerning the relationship of occupational exposure and smoking to chronic obstructive bronchopulmonary disease (cont.)*

Author, year, country, reference	Number and type of population	Results
Sluis-Cremer et al., 1967, South Africa (209). (cont.)	827 miners and nonminers over 35 years of age.	The dose relationship of cigarettes and COPD <sup>1</sup> symptoms was much more noticeable among those exposed to dust. The authors stressed the synergistic actions of cigarette smoking and dust exposure.
Bouhuys et al., 1969, U.S.A. (39).	455 male cotton textile workers (214 exposed to dust in carding and spinning rooms, 241 not exposed).	Those exposed to dust manifested a significantly greater prevalence of byssinosis symptoms than nonexposed. Smokers manifested a significantly greater prevalence of byssinosis symptoms than nonsmokers. No significant differences in Monday morning FEV <sub>1</sub> values were observed between smokers and nonsmokers. Prevalence of byssinosis symptoms did not show any relationship to length of employment.
Bouhuys et al., 1969, U.S.A. (38).	216 male hemp workers and 247 workers in other industries in same region, 20-69 years of age.	Hemp workers (especially the older ones) were noted to have different smoking habits from control group—fewer heavy smokers, more light smokers, more ex-smokers due to doctor's orders. Aged 20-49 — a. No difference in FEV <sub>1.0</sub> <sup>1</sup> values between controls and hemp workers in any smoking category. b. No difference in FEV <sub>1.0</sub> values between men in different smoking categories. Aged 50-69 — a. Hemp workers manifested decreased FEV <sub>1.0</sub> values in all smoking groups except for heaviest smokers. Ex-smokers had lowest FEV <sub>1.0</sub> values. b. Those smoking most had lower FEV <sub>1.0</sub> values as compared with light and non-smokers. The authors conclude that: There appears to be no synergism between smoking and hemp exposure as to effect on FEV <sub>1.0</sub> although the selection process whereby those with symptoms have a greater tendency to stop smoking may obscure such a relationship.
Chester et al., 1969, U.S.A. (49).	139 male chlorine plant workers (55 with history of severe ex- posure).	Chlorine-exposed group manifested no difference in symptoms and a decreased MBC value when compared with non-exposed group. Smokers in chlorine-exposed group had significantly decreased MBC and FEV values as compared with non-smokers in non-exposed group.
Greenberg et al., 1970, England (97).	121 workers in washing powder factory (48 found to be sensitized to product, 73 not).	Sensitized group manifested lower FEV <sub>1.0</sub> /FVC <sup>1</sup> values as compared with nonsensitized group even after smoking habits were controlled for.
Tokuhashi et al., 1970, U.S.A. (218).	801 male miners	Increased mine exposure was associated with residual volume and FEV abnormalities even after adjustments for age and smoking. A systematic exposure-impairment relationship was noted only among smokers while relatively few nonsmokers showed COPD impairment. Smoking miners manifested more X-ray alterations and COPD symptoms than nonsmokers, regardless of number of years of underground exposure.

<sup>1</sup> See Glossary of Terms in Bronchopulmonary table A4.

TABLE A10.—*Experiments concerning the effect of the chronic inhalation of NO<sub>2</sub> upon the tracheobronchial tree and pulmonary parenchyma of animals*

Author, year, country, reference	Animal	Results
Freeman and Haydon, 1964 U.S.A. (90).	Sprague-Dawley rats.	25 p.p.m.: (a) after 37-41 days—moderate hypertrophy and hyperplasia of bronchial and bronchiolar epithelium. (b) after 146-157 days—(1) Advanced hypertrophy and hyperplasia of bronchial and bronchiolar epithelium, (2) Increased lung volume. (3) Proliferation of connective tissue.
Haydon et al., 1965 U.S.A. (107).	Sprague-Dawley rats.	12.5 p.p.m. to death: (a) Hypertrophy and occasional metaplasia of bronchial and bronchiolar epithelium. (b) Increase in number of actively secreting goblet cells.
Haydon et al., 1967 U.S.A. (106).	Albino rabbits.	8-12 p.p.m. for 4 months: (a) Abnormal dilatation of peripheral air spaces. (b) Decreased density of alveolar walls. (c) Hypertrophy and hyperplasia of bronchial epithelium (especially terminal bronchiolar). (d) Increase in size of alveolar ducts. (e) Increased elastic tissue staining. (f) Increased alveolar size.
Freeman et al., 1968, U.S.A. (91).	Sprague-Dawley rats.	0.8 p.p.m.-2 p.p.m. for entire lifespan: (a) Alveolar distention. (b) Reduction in number of cilia. (c) Epithelial inactivity ("dormancy").
Freeman et al., 1968, U.S.A. (89).	Sprague-Dawley rats.	18 p.p.m. (a) 5 days—terminal bronchiolar epithelial hypertrophy. (b) 4 weeks—(1) Widespread bronchiolar epithelial hypertrophy. (2) Non-necrotizing emphysema.
Blair et al., 1969, U.S.A. (32).	Female Swiss Albino mice.	0.5 p.p.m.: (a) 6 hours/day for 3 months—pneumonitis. (b) 24 hours/day for 3 months—(1) Respiratory bronchiolar obstruction. (2) Alveolar expansion and bronchiolar inflammation consistent with early focal emphysema.
Kleinerman, 1970, U.S.A. (136).	Male Syrian Golden hamsters.	100 p.p.m. for 5½ hours: (a) thymidine autoradiography—intense burst of proliferation of epithelium returning to normal in 4 days (more persistent distally). (b) electron microscope—(1) Decreased number of secretory cells + secretory granules. (2) Increased number of lysosomal structures. (3) No change in number of ciliated cells.

TABLE A13.—*Experiments concerning the effect of cigarette smoke or its constituents upon ciliary function*

Author, year, country, reference	System	Method <sup>1</sup>	Results
Mendenhall and Shreeve, 1937, U.S.A. (164).	<i>In vitro</i> : Calf trachea.	Cigarette smoke by direct appli- cation or in solution.	Controls—ciliary activity depressed approxi- mately 4 percent. Experimental—ciliary activity depressed ap- proximately 40 percent.
Rakieten et al., 1942, U.S.A. (188).	<i>In vitro</i> : (a) rabbit and rat tracheal rings. (b) human nasal mucous membrane	I. Nicotine in Locke-Ringers solution. II. Cigarette smoke in solution.	I. Ciliary activity depressed only upon ex- posure to 100 mg. percent solution. II. Ciliary activity depressed after 15-20 min- utes exposure depending on concentration of smoke in solution.
Kordik et al., 1952, England (137).	<i>In vitro</i> : Rabbit trachea	Nicotine in Locke's solution.	Nicotine at 10 <sup>-3</sup> g./cc had no effect on ciliary activity.
Hilding, 1956, U.S.A. (120).	<i>In vitro</i> : Cow trachea	Cigarette smoke (direct exposure).	All tracheas showed depressed or absent ciliary activity.
Krueger and Smith, 1958, U.S.A. (139).	<i>In vivo</i> : Rabbit trachea	Cigarette smoke.	Cigarette smoke decreased ciliary activity by approximately 200 beats/minute.
Dalhamn, 1959, Sweden (59).	<i>In vivo</i> : I. Rat trachea <i>In vitro</i> : II. Rabbit trachea III. Human ciliated mucosa	Cigarette smoke.	I. 7/10 showed cessation of ciliary activity after one exposure. II. 6/10 showed cessation of ciliary activity after one exposure. III. 6/7 showed cessation of ciliary activity after one cigarette exposure.
Falk et al., 1959 U.S.A. (80).	<i>In vitro</i> : Rat and rabbit tracheal epithelium.	Cigarette smoke.	Decreased ciliary activity noted on exposure to cigarette smoke: (a) Repetitive exposure was associated with persistence of response over longer periods of time. (b) "Tar"-rich cigarette was more inhibitory than "tar"-poor. (c) Filtered smoke was less inhibitory than unfiltered.
Ballenger, 1960, U.S.A. (25).	<i>In vitro</i> : Human bronchial and tracheal epithelium obtained during anesthesia.	Cigarette smoke in solution.	Ciliary activity was fully inhibited within 5-28 minutes of exposure depending upon concen- tration of smoke in solution.

TABLE A13.—*Experiments concerning the effect of cigarette smoke or its constituents upon ciliary function (cont.)*

Author, year, country, reference	System	Method <sup>1</sup>	Results
Wynder et al., 1963, U.S.A. (236).	<i>In vivo:</i> Fresh water mussel ciliated epithelium.	Cigarette smoke; and its fractions in solution.	Unfiltered cigarette smoke—ciliastasis by 2nd- 5th puff. Acid (phenolic) fraction solution—immediate ciliastasis. Whole extract fraction solution—no ciliastasis. Neutral fraction solution—no ciliastasis. 1 percent phenol solution—immediate ciliasta- sis.
Dalhamn and Rylander, 1964, Sweden (61).	<i>In vivo:</i> Cat trachea.	Cigarette smoke.	Unfiltered cigarettes—ciliastasis in 3/5 cats after 5 cigarettes. Filtered cigarettes—no ciliastasis after 8 ciga- rettes (5 cats). Controls—no ciliastasis (5 cats).
Ballenger et al., 1965, U.S.A. (26).	<i>In vitro:</i> Human ciliated tracheal epithelium obtained during anesthesia.	Nicotine in solution.	Initial stimulation of activity followed by de- cline and complete ciliastasis after 12–24 hours of exposure.
Dalhamn and Rylander, 1965, Sweden (62).	<i>In vivo:</i> Cat trachea.	Cigarette smoke.	The longer the time interval between expo- sures, the more puffs were required to cause ciliastasis.
Wynder et al., 1965, U.S.A. (235).	<i>In vivo:</i> Fresh water mussel ciliated epithelium	Various compounds in solution.	Formic, acetic, propionic, benzoic acids all more ciliatoxic than phenol. Oxalic acid less ciliatoxic than phenol. Formaldehyde, acrolein more ciliatoxic than phenol.
Carson et al., 1966, U.S.A. (44).	<i>In vivo:</i> Cat trachea.	Cigarette smoke.	<i>Percent decrease in ciliary activity</i> Control ..... 0 Unfiltered smoke ..... 53 Cellulose acetate filter ..... 46 Carbon cellulose acetate filter ..... 30
Dalhamn, 1966, Sweden (60).	<i>In vivo:</i> Cat trachea.	Cigarette smoke.	<i>Mean number of puffs required to produce ciliastasis</i> No filter ..... 91 Charcoal filter ..... 170 Commercial cellulose acetate filter ..... 194 Charcoal and acetate filter ..... 512 Cambridge filter ..... 600
Kensler and Battista, 1966, U.S.A. (135).	<i>In vivo:</i> Rabbit trachea, cat trachea, dog trachea, monkey trachea. rat trachea.	Cigarette smoke and components in Tyrode's solution.	Rabbit trachea—Total smoke condensate of 3 cigarettes, gas phase condensate of 7 ciga- rettes caused similar ciliastasis. Other species—All found sensitive to ciliastatic components of cigarette smoke. Bulk of ac- tivity noted in gas phase (HCH, formalde- hyde, acrolein).

TABLE A13.—*Experiments concerning the effect of cigarette smoke or its constituents upon ciliary function (cont.)*

Author, year, country, reference	System	Method <sup>1</sup>	Results
Dalhamn and Rylander, 1967, Sweden (63).	<i>In vivo:</i> Cat trachea.	Cellulose acetate- filter cigarettes with varying amounts of "tar" but simi- lar gas phases.	Increased amounts of tar were associated with decreased number of puffs required to inhibit ciliary activity.
Dalhamn and Rylander, 1968, Sweden (64).	<i>In vivo:</i> Cat trachea.	Unfiltered and Cambridge-filter cigarettes.	Whole smoke found to be markedly more toxic to ciliary activity than volatile (gas) phase at lower dosages (puff volume). This differ- ence diminishes with increasing puff volume.
Kaminski et al., 1968, U.S.A. (133).	<i>In vivo:</i> Cat trachea.	Whole and filtered cigarette smoke exposed or unex- posed to "wet chamber" made to stimulate oral mucosa and saliva.	Wet chamber adsorption significantly reduced the ciliastatic activity of whole smoke, but did not affect the ciliastatic activity of smoke previously filtered by Cambridge or charcoal filters.
Krahl and Bulmash, 1969, U.S.A. (138).	<i>In vivo:</i> Common mollusk ciliated epithelium.	Cigarette smoke dissolved in sea water.	Significant ciliastasis, reversible.
Battista and Kensler, 1970, U.S.A. (28).	<i>In vitro:</i> Chicken tracheal epithelium.	Cigarette smoke or HCN in Tyrode's solution.	The authors observed that: (1) The more diluted smoke required more puffs to cause ciliastasis. (2) Activated charcoal filtered smoke was less ciliastatic than cellulose acetate filtered smoke and also contained less HCN and acrolein. (3) HCN alone was ciliastatic but recovery was more rapid than after cigarette smoke alone. They conclude that the gas phase components are more related to ciliastasis (as particulate matter is not significantly decreased by char- coal filtration while HCN and acrolein are).
Battista and Kensler, 1970, U.S.A. (29).	<i>In vivo:</i> Hen trachea.	Cigarette smoke.	The authors observed that: (1) Whole smoke acutely depressed ciliary activity in 4-6 puffs. (2) Gas phase was only slightly less depres- sant than whole smoke. (3) Chronic exposure (1 cigarette/day for 32 days) to smoke resulted in no apparent permanent defect in ciliary activity (al- though mucous production was signifi- cantly increased).

TABLE A13.—*Experiments concerning the effect of cigarette smoke or its constituents upon ciliary function (cont.)*

Author, year, country, reference	System	Method <sup>1</sup>	Results
Dalhamn and Rylander, 1970, Sweden (65).	<i>In vivo</i> : Cat trachea.	Unfiltered cigarette and cigar smoke.	Average number of puffs required to arrest ciliary activity Cigarette smoke . . . . . 73 } (p<0.01) Cigar smoke . . . . . 114 } The authors note that cigar smoke is of a different pH and that it contains more iso- prene, acetone, toluene, and acetonitrile.
Kennedy and Elliott, 1970, U.S.A. (134).	<i>In vivo</i> : Protozoan (ciliated).	Mainstream cigarette smoke.	Electron microscopic observations: (1) After 7 minutes exposure—alteration of mitochondrial structure. (2) After 42 minutes exposure—destruction of internal mitochondrial membrane struc- ture. (3) Gas phase alone, while ciliatotoxic, did cause mitochondrial swelling but no dis- ruption of membrane structure.

<sup>1</sup> Unless otherwise stated, method entailed the direct observation of ciliary activity using markers.



TABLE A14.—Experiments concerning the effect of cigarette smoke on pulmonary surfactant and surface tension

Author, year, country, reference	System	Method	Results			
Miller and Bondurant, 1962, U.S.A. (165)	Rat lung extracts	Cigarette smoke: (1) Applied to extract. (2) Exposure of rats.	(1) Exposure to cigarette smoke was associated with decreased surface tension in lung extract. (2) Surface tension of rats (lung extracts) exposed to cigarette smoke was decreased as compared with those not exposed.			
Cook and Webb 1966, U.S.A. (57)	40 subjects undergoing bronchoscopy: 14 normal 7 nonsmokers with pulmonary disease 19 smokers with and without pulmonary disease.		Surface tension values of surfactant		Stability index (reflects surfactant activity)	† Values significantly different from values of normals at $p < 0.02$ level.
			20 percent area	100 percent area		
		Normal .....	6.5	60.0	1.61	
		Pulmonary patients .....	†17.0	†50.0	1.00	
		Chronic smokers ..	15.7	51.0	1.04	
Giammona 1967, U.S.A. (94)	<i>In vitro</i> : Surfactant material induced from dogs and rats.  <i>In vivo</i> : Dogs, cats, and guinea pigs.	Exposed to cigarette smoke for 3 hours/day for up to 3 weeks.	<i>In vitro</i> : Exposure to cigarette smoke was associated with a significant decrease in maximal surface tension.  <i>In vivo</i> : Dogs and cats (exposed for 1 week)—no significant change. Guinea pigs (exposed for 3 weeks)—significant decrease in maximal surface tension.			
Webb, et al. 1967, U.S.A. (224)	Bronchial washing, from dog lungs.	Direct exposure to cigarette smoke.	Surface tension values of surfactant		Stability index	
			Number	20 percent area	100 percent area	
		Control .....	11	7.1	60.7	1.60
		Smoke .....	10	18.7	45.8	0.84